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	/ZV/		repe	Apostol et al., "A cell-based assay for aggregation inhibitors as therapeutics of polyglutamine- repeat disease and validation in <i>Drosophila</i> "; PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES" 2003 May 13, 100 (10):5950-5955.							
	/ZV/			Swanson et al., "Methionine Sulfoximine Reduces Cortical Infarct Size in Rats After Middle Cerebral Artery Occlusion", STROKE, vol. 21, No. 2, February 1990, pp 322-327.							
	IZVI			Lamar et al., "The duration of the inhibition of glutamine synthetase by methionine sulfoximine", BIOCHEMICAL PHARMACOLOGY, vol. 17, pp 636-640, 1968							
	/ZV/		hype	Takahashi et al., "Inhibition of brain glutamine accumulation prevents cerebral edema in hyperammonemic rats", THE AMERICAN PHYSIOLOGICAL SOCIETY, 1991, pgs. H825-H829.							
	/ZV/		Hack et al., "Glufosinate Ammonium-Some aspects of its mode of action in mammals", FD CHEM.TOXIC., vol. 32, no. 5, pp 461-470, 1994.								
	Gill et al., "The crystal structure of phosphinothricin in the active site of glutamine synthetase Illuminates the Mechanism of Enzymatic Inhibition", BIOCHEMISTRY 2001, 40, 1903-1912.										
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IZVI			Hertz et al., "Astrocytes:Glutamate Producers for Neurons", JOURNAL OF NEUROSCIENCE RESEARCH, 57:417-428 (1999).							
IZVI	·	repe	Apostol et al., "A cell-based assay for aggregation inhibitors as therapeutics of polyglutamine- repeat disease and validation in <i>Drosophila</i> ", PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES" 2003 May 13, 100 (10):5950-5955.							
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/ZV/	AA	Blei, Andres T., et al., "Ammonia-Induced Brain Edema and Intracranial Hypertension in Rats After Portacaval Anastomosis," <i>Hepatology</i> 19(6): 1437-1444, June 1994.						
	АВ			Errors of Urea Synthesis," <u>I</u> Pediatrics. 5: 140-165, Lon	n: Scriver CR, Lloyd Jk, eds. Genetic don: Butterworths, 1985.			
	AC	Brusilow, Saul W., et al., "Urea Cycle Disorders: Diagnosis, Pathophysiology, and Therapy," Advances in Pediatrics 43:127-170, 1996.						
	AD		Butterworth, R.F., "Effects of Hyperammonaemia on Brain Function," <i>J. Inher. Metab. Dis.</i> 21(1):6-20, 1998.					
	AE	Cordoba, Juan, et al., "Brain Edema and Hepatic Encephalopathy," Seminars in Liver Disease 16(3): 271-280, 1996.						
	AF	Folbergrova, J., "Free Glutamine Level in the Rat Brain In Vivo After Methionine Sulphoximine Administration," <i>Physiologia Bohemoslovenica</i> 13:21-26, 1963.						
	AG	Gershoff, S.N., et al.,"The Relative Effect of Methionine Sulfoximine on Different Animal Species," J. Nutr. 45:451-458, 1951.						
	АН	Häussinger, Dieter, et al.,"Pathogenesis of Hepatic Encephalopathy," Journal of Gastroenterology and Hepatology 17:S256-S259, 2002.						
	Al	Hawkins, Richard, et al., "Effect of Reducing Brain Glutamine Synthesis on Metabolic Symptoms of Hepatic Encephalopathy," <i>Journal of Neurochemistry</i> 60(3):1000-1006, 1993.						
	AJ	Hawkins, Richard, et al., "Hyperammonaemia Does Not Impair Brain Function in the Absence of Net Glutamine Synthesis," <i>Biochem. J.</i> 277:697-703, 1991.						
	AK	Hirata, Takahiko, et al., "Impaired Pial Arteriolar Reactivity to Hypercapnia During Hyperammonemia Depends on Glutamine Synthesis," Stroke 27(4): 729-736, 1996.						
	AL				vents the Accumulation of Large Neutral Surgical Research 36:349-353, 1984.			
	АМ	Krakoff, Irwin Ther. 2:599-6		Effect of Methionine Sulfoxim	ine in Man," <i>J. Pharm. Experimenetal</i>			
	AN			ration of the Inhibition of Glul Pharmacology 17:636-642,	itamine Synthetase by Methionine 1968.			

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/Z\	V/	AO	Master, Sonali, et al., "Cerebral Blood Flow and the Development of Ammonia-Induced Brain Edema in Rats After Portacaval Anastomosis," <i>Hepatology</i> 30(4): 876-880, 1999.							
AP		AP	Norenberg, Michael D., et al,"Fine Structural Localization of Glutamine Synthetase in Astrocytes of Rat Brain," <i>Brain Research</i> 161:303-310, 1979.							
		AQ		"Inhibition of γ-Glutamylcystein Synthetase by L-Methionine-S- al Chemistry 248(19): 6684-6690, 1973.						
		AR			-S-Sulfoximine as the Convulsant e National Academy of Sciences 66(2):					
		AS	Sellinger, Otto Z., et al., "Methionine Sulfoximine Seizures. VII. the Dissociation of the Convulsant and Glutamine Synthetase Inhibitory Effects," J. Pharmacology & Experime Therapeutics 161(1): 212-222, 1968.							
		АТ	Attenuates Increased Ext	al., "Methionine Sulfoximine, A Glutamine Synthetase Inhibitor, tracellular Potassium Activity During Acute Hyperammonemia," d Flow & Metabolism, 17:44-49, 1997.						
		AU		"Inhibition of Brain Glutamine Accumulation Prevents Cerebral emic Rats," <i>American Physiological Society</i> 261:H825-H829, 1991.						
		AV		cute Hyperammonemia in the Young Primate: Physiologic and tes," <i>Pediatric Research</i> 17(12):970-975, 1983.						
				ne Susceptibilty of Auditory Stimuli of Animals Treated with Experimental Neurology 15:157-165, 196.						
		AX		al., "Effect of an Inhibitor of Glutamine Synthesis (Methionine lia Toxicity and Metabolism," <i>J. Lab. & Clin. Med.</i> 64(3): 442-449,						
		AY	Watson, Alan J., et al. "Ti 1274, December 7, 1985.	Fransient Idiopathic Hyperammonaemia in Adults," <i>The Lancet</i> 1271-5.						
		AZ		I., "Inhibition of Glutamine Synthetase Reduces Ammonia-Induced at," Neuroscience 71(2): 589-599, 1996.						
		ВА		d Detoxification Processes in	roscopy Studies on NH4CI-Induced n Primary Astrocytes and Glioma Cells,					
		вв	Apostolakis et al., Brain F	Research Bulletin, Vol. 23, pp. 257-262 (1989).						
		вс	Ginefri-Gayet et al., Phar	macology Biochemistry and Behavior, Vol. 43, pp. 173-179 (1992).						
BD Takahashi et al., Circulati				ation Research, 71(5), 1220-1230 (November, 1992).						
	V	BE	Harth et al., J. Exp. Med.	, 189(9), 1425-1435 (May 3,	1999).					

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Appl. No.

: 10/758,415

Applicant

: William S. Brusilow

Filed

: January16, 2004

TC/A.U.

: 1614

Examiner

Docket No.

: 2930-109 Customer No.: 06449

Confirmation No.: 5654

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Director of the United States Patent and Trademark Office P.O. Box 1450 Alexandria, Virginia 22313-1450

Dear Sir:

During a review of the references filed on December 20, 2004, it was noted that the attached reference entitled "Urea Cycle Disorders: Diagnosis, Pathophysiology, and Therapy" was missing several pages. Attached is a complete copy of that reference for the Examiner's review.

Respectfully submitted,

Monica Chin Kitts

Attorney for Applicants Registration No. 36,105

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